

VOS-31

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Examiner : Not yet assigned  
Group : Not yet assigned  
Applicants : Leszek Wojnowski et al.  
Application No. : Not yet assigned  
Confirmation No. : Not yet assigned  
Filed : Concurrently herewith  
For : POLYMORPHISMS IN THE HUMAN hPXR GENE AND  
THEIR USE IN DIAGNOSTIC AND THERAPEUTIC  
APPLICATIONS

New York, New York  
March 8, 2002

Hon. Commissioner for Patents  
P.O. Box 2327  
Arlington, Virginia 22202

INFORMATION DISCLOSURE STATEMENT

Pursuant to 37 C.F.R. §§ 1.56 and 1.97, applicants, through their  
representatives, make of record the following documents\*:

Foreign Patent Documents

| <u>Country</u> | <u>Publication No.</u> | <u>Publication Date</u> |
|----------------|------------------------|-------------------------|
| Europe         | EP 1 024 193           | August 2, 2000          |
| PCT            | WO 99/48915            | September 30, 1999      |

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\* For the convenience of the Examiner, a completed Form PTO-1449, listing these documents, is attached.

### Other Documents

Bertilsson, G. et al., "Identification Of A Human Nuclear Receptor Defines A New Signaling Pathway For CYP3A Induction," *Proc. Natl. Acad. Sci. USA*, 95:12208-12213 (1998).

Dotzlaw, H. et al., "The Human Organ Receptor PXR Messenger RNA IS Expressed In Both Normal And Neoplastic Breast Tissue," *Clinical Cancer Research* 5:2103-2107 (1999).

Jones, S. et al., "The Pregnane X receptor: A Promiscuous Xenobiotic Receptor That Has Diverged During Evolution," *Molecular Endocrinology*, 14:27-39 (2000).

Kliewer, S. et al., "An Orphan Nuclear Receptor Activated By Pregnanes Defines A Novel Steroid Signaling Pathway," *Cell* 92:73-82 (1998).

Lehmann, J. et al., "The Human Orphan Receptor PXR Is Activated By Compounds That Regulate *CYP3A4* Gene Expression And Cause Drug Interactions," *Journal of Clinical Investigation* 102:1016-1023 (1998).

Pascussi, J.-M. et al., "Evidence For The Presence Of A Functional Pregnane X Receptor Response Element In The *CYP3A7* Promoter Gene," *Biochemical and Biophysical Research Communications*, 260:377-381 (1999).

Zhang, H. et al., "Rat Pregnane X Receptor: Molecular Cloning, Tissue Distribution, and Xenobiotic Regulation," *Archives of Biochemistry and Biophysics* 368:14-22 (1999).

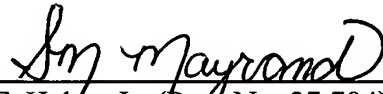
The documents listed above were cited in an International Search Report that issued in connection with International application No. PCT/EP00/08827 which is the parent international application of the above-identified application. For the Examiner's convenience, a copy of the Search Report is attached hereto. Applicants have not enclosed copies of these documents because they should have been supplied by the International Search Authority. However, if such copies have not been received, applicants will provide them.

This Statement is being submitted within three months of the application filing date. Accordingly, no payment of fees is required pursuant to 37 C.F.R.

§ 1.97(b)(1).

Applicants respectfully request that these documents be (1) considered by the Examiner prior to issuance of any patent from this application; and (2) printed on any patent that may issue from this application. Applicants also request that a copy of enclosed Form PTO-1449, as considered and initialed by the Examiner, be returned with the next communication.

Respectfully submitted,



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FORM PTO-1449

U.S. DEPARTMENT OF COMMERCE  
PATENT AND TRADEMARK OFFICEATTY. DOCKET NO.  
VOS-31SERIAL NO.  
Not yet assignedINFORMATION DISCLOSURE  
STATEMENT BY APPLICANTAPPLICANTS  
Leszek Wojnowski et al.CONFIRMATION NO.  
Not yet assignedFILING DATE  
Concurrently herewithGROUP  
Not yet assigned

## U.S. PATENT DOCUMENTS

| EXAMINER<br>INITIAL | DOCUMENT NUMBER | DATE | NAME | CLASS | SUBCLASS | FILING DATE<br>IF<br>APPROPRIATE |
|---------------------|-----------------|------|------|-------|----------|----------------------------------|
|                     |                 |      |      |       |          |                                  |
|                     |                 |      |      |       |          |                                  |

## FOREIGN PATENT DOCUMENTS

| EXAMINER<br>INITIAL | DOCUMENT NUMBER | DATE     | COUNTRY | CLASS | SUBCLASS | TRANSLATION |    |
|---------------------|-----------------|----------|---------|-------|----------|-------------|----|
|                     |                 |          |         |       |          | YES         | NO |
|                     | EP 1 024 193    | 08/02/00 | Europe  | C12N  | 15/60    |             |    |
|                     | WO 99/48915     | 09/30/99 | PCT     | C07K  | 14/00    |             |    |
|                     |                 |          |         |       |          |             |    |

## OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, Etc.)

| EXAMINER<br>INITIAL |  |
|---------------------|--|
|                     | Bertilsson, G. et al., "Identification Of A Human Nuclear Receptor Defines A New Signaling Pathway For CYP3A Induction," <i>Proc. Natl. Acad. Sci. USA</i> , 95:12208-12213 (1998).                                    |
|                     | Dotzlaw, H. et al., "The Human Organ Receptor PXR Messenger RNA IS Expressed In Both Normal And Neoplastic Breast Tissue," <i>Clinical Cancer Research</i> 5:2103-2107 (1999).   |
|                     | Jones, S. et al., "The Pregnane X Receptor: A Promiscuous Xenobiotic Receptor That Has Diverged During Evolution," <i>Molecular Endocrinology</i> , 14:27-39 (2000).   |
|                     | Kliwer, S. et al., "An Orphan Nuclear Receptor Activated By Pregnanes Defines A Novel Steroid Signaling Pathway," <i>Cell</i> 92:73-82 (1998).   |
|                     | Lehmann, J. et al., "The Human Orphan Receptor PXR Is Activated By Compounds That Regulate CYP3A4 Gene Expression And Cause Drug Interactions," <i>Journal of Clinical Investigation</i> 102:1016-1023 (1998).         |
|                     | Pascussi, J.-M. et al., "Evidence for the Presence of a Functional Pregnane X Receptor Response Element in the CYP3A7 Promoter Gene," <i>Biochemical and Biophysical Research Communications</i> , 260:377-381 (1999). |
|                     | Zhang, H. et al., "Rat Pregnane X Receptor: Molecular Cloning, Tissue Distribution, And Xenobiotic Regulation," <i>Archives of Biochemistry and Biophysics</i> 368:14-22 (1999).                                       |
|                     |  |

EXAMINER

DATE CONSIDERED

EXAMINER: Initial if citation considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not conformance and not considered. Include copy of this form with next communication to applicant.

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